THE COUNCIL FOR TOBACCO RESEARCH – U.S.A., INC. 110 EAST 50th STREET NEW YORK, N. Y. 10022 (212) 421-8885 Application for Research Grant (Use extra pages as needed) Date: June 23, 1975

1. Principal Investigator (give title and degrees):
Carlos W. M. Bedrossian, M.D.
Assistant Professor of Pathology, University of Texas Medical School at Houston, Assistant Pathologist and Director of Cytopathology, Hermann Hospital, Houston, Texas.

2. Institution & address:
The University of Texas Health Science
Center at Houston (Medical School)
P. O. Box 20036
Houston, Texas

3. Department(s) where research will be done or collaboration provided:

3. Department(s) where research will be done or collaboration provided:

Pathology Department

4. Short title of study:

Relationship Retween Dulmonary Emphysion and Liver Starbooks. Relationship Between Pulmonary Emphysema and Liver Cirrhosis on the Basis of an Acquired Deficiency of Alpha-1-Antitrypsin

- 5. Proposed storting date: January 1, 1976 6. Estimated time to complete: Two years
 7. Brief description of specific research aims:
- 7. Brief description of specific research aims:

The purpose of this investigation is to determine the incidence and pattern of the emphysema present in association with advanced cirrhosis of the liver in an autopsy population of smoking and non-smoking alcoholics. This approach will be a continuation of our interest in studying the background factors and determining the relationship of emphysema to other conditions as we have previously done with heart disease, systemic hypertension² and bronchogenic carcinoma³.

Brief statement of working hypothesis:
The reason to investigate the relationship between emphysema and cirrhosis is based on the hypothesis that the latter will result in an acquired deficiency of alpha-1-antitrypsin (A1AT). As it is known, the liver is the site of production of AlAT believed to be the principal inhibitor of the proteases4. Congenital deficiency of this enzyme was first associated with emphysema appearing in young persons. Subsequently, lack of AIAT has been linked to the development of cirrhosis in children⁶. More recently, either emphysema⁷ or cirrhosis⁸, or both9, have been reported in adults and elderly patients with AIAT deficiency. most instances, it has been postulated that the basic defect is the enzyme deficiency, whereas the disease processes represent secondary manifestations 10,11, However, the possibility of the cirrhosis being the primary defect with subsequent appearance of the enzyme deficiency and resulting emphysema has never been explored. The purpose of our study is to test this hypothesis by initially determining the extent of the association between cirrhosis and emphysema. ld .

9. Details of experimental design and procedures (append extra pages as necessary) Lungs from consecutive autopsies of patients with cirrhosis will be included in the study. After careful removal, inflation and fixation will be accomplished with formaldehyde vapor by means of a modified Weibel's apparatus 12. See 1997 to the the programme to the control of the

Low intensity roentgenograms will be performed on high contrast industrial films in a Faxitron machine after which tantalum bronchograms will be performed. Lungs will then be cut parasagitally at 2 cm. intervals with an electric knife. Sections will be examined with a zoom stereoscopic dissection microscope and interesting areas will be photographed.

In the sections of inflated specimens it will be possible to classify emphysema anatomically as centrilobular, panlobular, paraseptal, and irregular, according to the criteria of the American Thoracic Society 13. A detailed assessment of each lung will determine certain parameters of the emphysema present such as distribution of the process within the lung, localization of the lesions within the secondary lobules and degree of pulmonary involvement.

San Commence of the San Commence Hepatic cirrhosis will be studied pathologically in the same autopsied patients by gross and histopathological techniques to include H and E and other stains such as Masson's trichrome, P.A.S., Snook's reticulin and others. Cirrhosis of the liver will be classified according to the criteria set forth by Rubin and Popper as: regular, mainly monolobular cirrhosis (Laennec's or nutritional cirrhosis) and irregular, mainly multilobular cirrhosis (postnecrotic or posthepatitic cirrhosis) 17. The incidence of emphysema among all cirrhotic patients in general and the specific types of cirrhosis will then be statistically determined. Correlation will be made between the different types and degrees of emphysema associated with the various types of liver cirrhosis taking into account the smoking habits of the patients. Special attention will be paid to comparing the various specimen x-rays with the in vivo chest roentgenograms in an attempt to determine if the emphysema present could be diagnosed radiographically. In <u>vivo</u> and postmortem serum trypsin inhibitory capacity (S.T.I.C.) will be determined according to Hammarsten in the same patients whose lungs and liver are included in the study 15 . The S.T.I.C. values will then be correlated with the respective degrees of emphysema and cirrhosis. In addition, all sera will be phenotyped as to the exact level of A1AT. ATTAL AREA THE TOTAL

10. Space and facilities available (when elsewhere than item 2 indicates, state location):

For the study of lung and liver tissues the project will have the support of the Histopathology Laboratory of the Pathology Service at Hermann Hospital. For the special study of inflated and fixed lung specimens floor space will be made available in the Department of Pathology of the University of Texas Medical School at Houston, Texas. Facilities will also be provided by the Department of Pathology for the study of the in vivo chest roentgenograms and the various specimen Xrays. Serum trypsin inhibitory capacity determinations will be done at the Clinical Pathology Laboratory of Hermann Hospital.

Dr. Warren C. Miller and Dr. Donald C. Cannon are co-investigators in the project. Dr. Warren C. Miller will cooperate with us in identifying patients to enter the study. Dr. Donald C. Cannon will cooperate with us in determination of serum trypsin inhibitory capacity and phenotyping of the sera.

11. Additional facilities required:

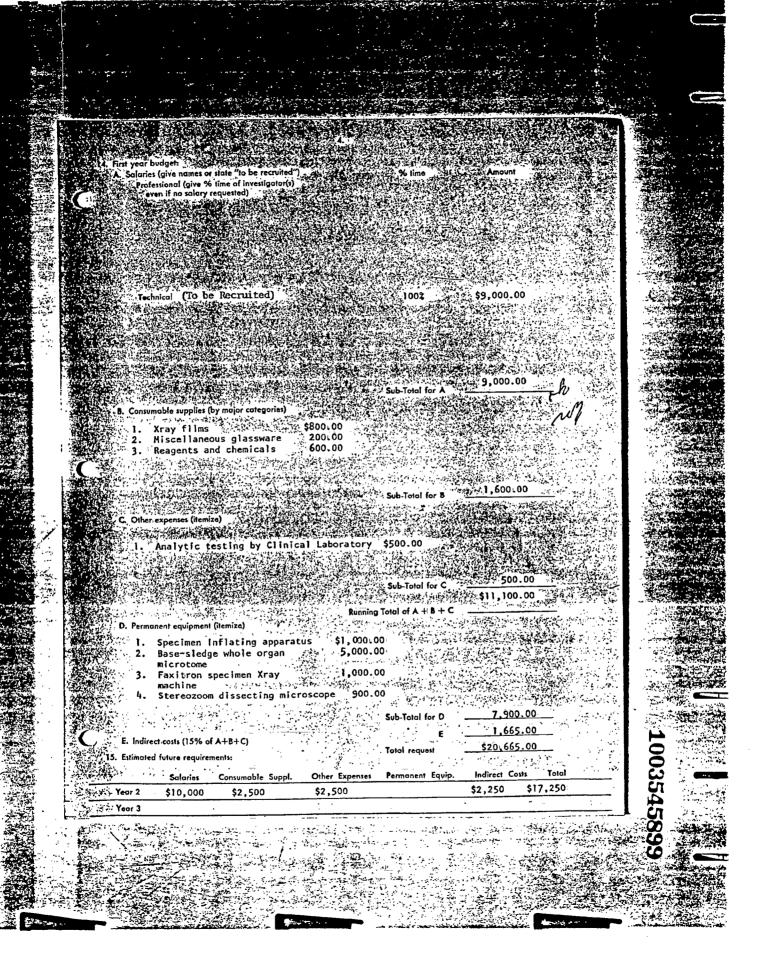
The proposed program does not require collaboration with other institutions. Nevertheless, potential cooperation is available from the institutions where the principal investigator has previously been trained. Specifically, a large series of emphysematous lungs studied by the whole organ section technique is available to us at a Research Center in Florida. The incidence and patterns of cirrhosis could easily be determined among these emphysematous cases.

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12. Biographical sketches of investigator(s) and other professional personnel (append):

13. Publications: (five most recent and pertinent of investigator(s); append list, and provide reprints if available).

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34-12	Other sources of financial support				
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- REFERENCES:

 1. Foraker, A. G., Bedrossian, C.W.M., and Anderson, A.E. Jr.: Myocardial dimensions and proportions in pulmonary emphysema. Arch. Pathol. 90:344-347, 1970.
- Anderson, A.E., Jr., Bedrossian, C.W.M. and Foraker, A.G.: Systemic blood pressure in subjects with and without emphysema. A mer. Rev. Resp.
- Bedrossian, C.W.M.: Coexistent emphysema and bronchogenic carcinoma.
- Cancer Bulletin 24:102-107, 1972.

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 Human Genetics 6:207, 1968.

 Laurell, C.B. and Eriksson, S.: The electrophoretic alpha-1 globulin
- pattern of serum in alpha-l-antitrypsin deficiency. Scand. J. Clin. Lab. Invest. 15:132, 1963.
 Sharp, H.L., Bridges, R.A., Krivit, W. and Freier, E.F.: Cirrhosis
- associated with alpha-l-antitrypsin deficiency: A previously unrecognized inherited disorder. J. Lab. Clin.:Med. 73:934-939, 1969. (多数)。
- Tarkoff, M.P., Kueppers, F. and Miller, W.F.: Pulmonary emphysema and alpha-l-antitrypsin deficiency. Amer. J. Med. 45:220-228, 1969.
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- Gherardi, G.J.: Alpha-1-antitrypsin deficiency and its effect on the liver. Human Pathol. 2:173-175, 1971.
- Eriksson, S.: Studies_in alpha-l-antitrypsin deficiency. Acta Med. Scand. 177, Suppl. 432, 1965. Sharp, H. A.: Alpha-1-antitrypsin deficiency. Hosp. Pract. 83, 1971.
- 11.
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 - American Thoracic Society: chronic bronchitis, asthma, and pulmonary emphysema: A statement by the Committee on Diagnostic Standards for nontuberculous respiratory diseases. Amer. Rev. Res. Dis.
 - 85:762, 1962.
 Rubin, E. and Popper, H.: The evolution of human cirrhosis as deduced from observations in experimental animals. Med. 46:163, 1967.

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PLACE AND DATE OF BIRTH:

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CHILDREN:

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EDUCATION:

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M.D., University of Sao Paulo, Ribeirao Preto Medical School, Ribeirao Preto, Sao Paulo, Brazil.

Lieutenant, Medical Corps, Brazilian Air Force, 1968.

POST GRADUATE ACTIVITIES:

1968-1969, Chief of Laboratories, 4th Aerial Zone Hospital Brazilian Air Force, Sao Paulo, Brazil, Volunteer Assis ant, Departments of Pathology, Santa Casa, Paulista de Medicina and Juqueri Hospitals, Sao Paulo, Brazil.

1969-1971, Resident in Pathology and Research Fellow in Pulmonary Pathology, Baptist Memorial Hospital, 🎇 Jacksonville, Florida, U.S.A.

建一个"这个"。 1971-1972, Fellow in Anatomical Pathology, The University of Texas M.D. Anderson Hospital and Tumor Institute, Texas Medical Center, Houston, Texas, U.S.A.

1972-1973, Fellow in Diagnostic Cytopathology and Pulmonary Pathology, Baylor College of Medicine, and Chief Resident in Pathology, Ben Taub General Hospital Texas Medical Center, Houston, Texas, U.S.A.

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HOSPITAL APPOINTMENTS:	1973-1975: Assistant Pathologi	st and Director of Cyto-
	pathology, University Hospital,	
	Hospital, and Oklahoma Gbildren	's Memorial Hospital,
	The University of Oklahoma Heal	th Sciences Center,
Televida Arena (da100 Erana	Oklahoma City, Oklahoma.	
A STATE OF THE STA	1975-Present: Assistant Pathol	
	pathology, Hermann Hospital, Te	
	Texas I f f f f f f f f f f f f f f f f f f	建分子的 医神经神经病 医乳毒素
ACADEMIC POSITIONS:	1973-1975: Assistant Professor	of Pathology Colleges of
	Medicine and Dentistry, The Uni	
	Sciences Center, Oklahoma City,	
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	Associate Professor and Character technology, College of Allied H	
	University of Oklahoma Health S	
	City, Oklahoma	A Company of the Comp
		Cor of Pathology University
	1975-Present: Assistant Profes of Texas Medical School at Hous	
1	Houston, Texas	
LICENSES AND	1968 - Conselho Regional de Me	dicine, Sao Paulo, 💛 🚎
CERTIFICATES: - XX	Brazil, License #13110	
	1970 - Educational Council for	
	Graduates, Certificate ; 1971 - State of Texas Board of	
	Basic Sciences, Certific	
	1972 - Texas State Board of Med	
Also the state of	License #D-9356	Entropy (April 1997) Herring Color (April 1997)
	1973 - Oklahoma State Board of	
	11/4 License #10019 11/4/14	
	1974 - American Board of Patho	
	in Anatomic and Clinica	l Pathology
PROFESSIONAL	Service of the servic	A STATE OF THE STA
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FIELDS OF SCIENTIFIC	Diagnostic Cytopathology	
INTEREST:	tlinical-pathologic correlation	ns in chronic obstructive
	pulmonary disease.	. • •
	Examination of excised lungs by	pathological and
	radiological methods.	ومحجدة المحاجد في
	Electron microscopy of cancer a	ing other pulmonary

HONORS AND AWARDS: 🎋 Service of the Company of the Compan

1966, Academic Merit Diploma, The University of Sao Paulo, Ribeirao Preto, Sao Paulo, Brazil.

1967, Best Student in the Basic Sciences Award, The University of Sao Paulo, Ribeirao Preto, Sao Paulo,

1972, Annual Clinical Training Research Project Award, The University of Texas M.D. Anderson Hospital and Tumor Institute, Houston, Texas.

1974, "Who's Who in Texas", United States Public Relations Service, Atlanta, Georgia.

ELECTED OFFICE.

Branch Branch Branch Control

POST GRADUATE STUDIES:

President, Oklahoma Society of Cytopathology. 1974-75.
Pathology of the Lung. University of California, San Diego, La Jolla, California. Averyll A. Liebow, Director, June 23-28, 1969.

Gynecologic and Obstetric Pathology, Armed Forces Institute of Pathology, Washington, D.C., Henry J. Norris, Director, October 26-30, 1970.

Dermal Pathology. The University of Texas M. D.

·--

Anderson Hospital and Tumor Institute, Houston, Texas. J. Leslie Smith, Jr., Director, August 23-27, 1971.

Clinical Pathology of Respiratory Diseases. The Association of Clinical Scientists, Chicago, Illinois. William Sunderman, Director, November 3-5, 1972.

A Symposium on Diseases of the Chest, The Fleischner Society, Boston, Massachusetts, Leo G. Rigler, Direct May 26-29, 1973.

PRESENTATIONS:

- Pulmonary Emphysema" to Florida Academy of Sciences, Jacksonville University, Jacksonville, Florida, March 5, 1970.
- 2. "Bronchitis Cronica y Enfisema Pulmonar" to the Mexican Patholgists Association, Sequros Sociales Hospital, Mexico City, Mexico, June 19, 1970.
- 3. "An Eclectic Approach to Lung Morphology" to the Brazilian Pathologists Association, Northeast Chapter, Recife, Pernamuco, Brazil, January 14, 1972.
- 4. "Interstitial Pneumonitis Associated with Bleomycin Therapy" to the International Academy of Pathology, Cincinnati, Ohio, March 15, 1972.
- 5. "Lung Cancer and Emphysema" to the Texas Chapter American College of Chest Physicians, San Antonio, Texas, May 13, 1972.
- 6. "Bleomycin Pulmonary Toxicity In Clinical and Experimental Subjects" to the Medical Research Division, Bristol Laboratories, Syracuse, N.Y., May 25, 1973.
- 7. "Ultrastructure of the Lung in Loeffler's Pneumonitis" to the Latin-American Pathologists Association,
 Merida, Yucatan, Mexico, November 27, 1973.
- 8. "The Lung in Cystic Fibrosis of the Pancreas" to the Latin-American Pathologists Association, Merida, Yucatan, Mexico, November 27, 1973.
- 9. "Electron Microscopic Changes and some Experimental Aspects of Bleomycin Pulmonary Toxicity" to the National Cancer Institute, Division of Cancer Treatment, NIH, Bethesda, Maryland, February 15, 1974.
- 10. "Esophageal Cytopathology: Review of Experience and Presentation of Interesting Cases" to the Texas Society of Cytology, Dallas, Texas, April 27, 1974.
- 11. "The Role of Cytopathology in the Diagnosis of Gastroesophageal Diseases" to the Oklahoma Society of Cytopathology, Oklahoma City, Oklahoma, May 15, 1974.
- 12. "Experimental Bleomycin Pulmonary Toxicity" to the Brazilian Society of Pathology, Curitiba, Parana, Brazil, September 4, 1974.

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- PRESENTATIONS:

 13. "Ultrastructure of Human Bronchiolo-alveolar (continued)

 Carcinoma" to the Brazilian Society of Pathology,
 Curitiba, Parana, Brazil, September 7, 1974.

 14. "Pitfalls in Diagnosing, Cell Typing and Judging
 - Degree of Differentiation of Bronchogenic Carcinoma in Brush Cytology Specimens" to the
 American Society of Cytology, New York, New
 York, November 8, 1974.

- 1970: 1. Bohm, G.N., Bedrossian, C.W.M., De Luca, F. and Kerr, W.: The failure of nucleic acids to prevent the destruction of neurons in Chagas disease Rev. Lat. Amer. Pathol. 9:17-23 (Mexico).
- 2. Foraker, A.D., Bedrossian, C.W.M., and Anderson, A.E.: Myocardial dimensions and proportions in pulmonary emphysema. Arch. Pathol. 90:344-347.
- Bedrossian C W M 1971: 3. Anderson, A.E., Bedrossian, C.V.M., and Foraker, A.G.: Systemic blood pressure in subjects with and without emphysema. Amer. Rev. Resp. Dis. 103:576-578.

 4. Bedrossian, C.W.M., Anderson, A.E. and Foraker, A.G.: Comparison of methods
 - for quantitating bronchial morphology. Thorax 24:406-408, (London).
- 1972: 5. Luna, M.A., Bedrossian, C.W.M., Lichtiger, B. and Salem, P.: Interstitial pneumonitis associated with Bleomycin therapy. Amer. J. Clin. Pathol. 58:501-510.
 - 6. Bedrossian, C.W.M.: Coexistent emphysema and bronchogenic carcinoma. Cancer Bulletin 24:102-107.
- Bedrossian, C.W.M. and Martin, J.E.: Xeroradiography of the lung. Radiol. 107:217-218.
 - Bedrossian, C.W.M., Greenberg, S.D. and Duran, B.S.: Bronchial gland measurements: A continuing search for a yardstick. J. Exp. & Molec. Pathol. 18:219-224,
 - Bedrossian, C.W.M., Luna, M.D., MacKay, B. and Lichtiger, B.: Ultrastructure of pulmonary Bleomycin toxicity, Cancer 32:44-51.
 - 10. Wurlitzer, F., Bedrossian, C.W.M., Ayala, A.G. and McBride, C.: Problems in diagnosing and treating infiltrating lipomas. Amer. Surg. 39:240-243.
- 1974: 11. Stork, W.J., Greenberg, S.D. and Bedrossian, C.W.M.: Fatal sarcoidosis.
 VI Int. Conf. on Sarcoidoses, Tokio, Japan. pp 462-472, Univ. Park Press.
 - Bedrossian, C.W.M. Ultrastructural changes and some experimental aspects of Bleomycin pulmonary toxicity. In "New Drug Seminar" sponsored by National Cancer Institute, Division of Cancer Treatment, Bethesda, Md., p. 169, Feb.
- **多沙 13.** Bedrossian, C.W.M.: Current data regarding Bleomycin-induced pulmonary toxicity: An Audio Review. Intramed. Communications. In New York, N.Y., Vol. 2, Side 2, April.
- Bedrossian, C.W.M., Greenberg, S.D., and Williams, L.J.: Ultrastructure of the lung in Loeffler's pneumonitis. Amer. J. Med., 58:438-443.
 - Bedrossian, C.W.M., Greenberg, S.D., Singer, D., Jenson, J., and Rosenberg, H.: The lung in cystic fibrosis of the pancreas. Human Pathology (In press).

- 16. Weilbaecher, D.G., Bedrossian, C.W.M., Greenberg, S.D. and Bentick continued ____D.C.: Ultrastructure of human bronchioloalveolar carcinoma.

 Cancer (In press). D.C.: Ultrastructure of human bronchioloalveolar carcinoma.
 - 17. Bedrossian, C.W.M. and Rybka, D. L.: Bronchial brushing during fiberoptic bronchoscopy for the cytodiagnosis of lung cancer. Comparison with sputum and bronchial washings. Acta Cytologica. (Submitted for publication).
 - Bedrossian, C.W.M., Yawn, D.H., Greenberg, S.D. and O'Neal, R.M.: Experimentally-induced Bleomycin pulmonary toxicity in the pheasant - Part I: Histopathological findings. Lab. Invest. (Submitted for publication).
 - Bedrossian, C.W.M., Greenberg, S.D., Yawm, D.H. and O'Neal, R.M.: the angle Experimentally-induced Bleomycin pulmonary toxicity in the pheasant - Part II: Ultrastructural observations. Lab. Invest. (Submitted for publication.)

1975 Work in **Progress**

Bedrossian, C.W.M., Greenberg, S.D., Luna, M.A., Skinner, F.M. and Jenkins, D. E.: Pathological patterns of chronic obstructive pulmonary disease associated with "environmental" and "endogenous" The property of the second of the second filung cancer. (In preparation).

Home Phone
Date of Birth
Place of Birth Place of Birth Citizenship
Marital Status
Spouse's First Name
Number of Children
Social Security Number

Honors and Awards

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A. Personal Information:

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Number of Children
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Education

High School

High School

College or University

Medical School

Medical School

Medical School

Internship
Residencies

Internships
Residencies

Fellowships

Washington University School of Medicine,
St. Louis,
University of Chicago Medical School,
M.D.,
UCLA Hospital, 1960-61, Pathology
University of Chicago Clinics,
July 1961-June, 1964-Pathology
Summer Fellow in Pathology, 1958,
Washington University
U.S.P.H.S. Pre-doctoral Trainee, 1960, University of Chicago, Immunopathology U.S.P.H.S. Post-doctoral Trainee, 1961-1963, University of Chicago, Immunopathology Harvard College Scholarship, 1952-56 Phi Beta Kappa, Harvard College, 1956 Jackson Johnson National Scholarship, Washington University, 1956-1958 President, first year medical class, Washington University

Chicago University Scholarship, 1959-1960
Alpha Omega Alpha, University of Chicago
Sigma Xi, University of Chicago
Markle Scholar in Academic Medicine,
1967-1968

Award of Merit for Distinguished Service in Advancing the Cause of Blood Banking,
American Association of Blood Banks, 1967
Missouri, 1962 (certificate 28946)
New York, 1965 (certificate 95632)
California, 1970 (certificate C-32725)
American Board of Pathology (clinical and anatomic), 1967

Licensure ;

Board Certification

C. Professional Background:

Academic Appointments

Instructor of Pathology, University of Chicago, 1963-64
Instructor of Clinical Bacteriology and Pathology, University of
North Carolina, 1964-65

Assistant Professor of Pathology, Upstate Medical Center, 1965-68
Associate Professor (in absentia), Upstate Medical Center, 1969
Visiting Professor of Pathology, The University of New Mexico,
1972-present

Associate Clinical Professor of Pathology, University of Southern California, 1973-present

Specific teaching responsibilities

Autopsy case study for sophomores, 1960-61

Teaching in general histopathology for sophomores, 1961-62

Lectures in liver pathology for sophomores and in endocrine pathology for seniors, 1963-64

Lectures in general pathology and autopsy case study for sophomores. Lectures and laboratory training in clinical bacteriology, autopsy and surgical pathology training for pathology resident physicians, 1964-65

Lectures in immunology, blood banking, and clinical chemistry; conference leader in general pathology and clinical pathology; and gross autopsy review for sophomores. Lectures to medical technology students and blood bank technology students. Clinical pathology training of pathology resident physicians. 1965-68.

Specific teaching responsibilities

Teaching in blood banking and immunology, Foundation for Advanced Education in the Sciences, National Institutes of Health, 1968-69 Director of Resident Training, Clinical Pathology Department, NIH, 1969-70

Lectures and conference teaching in acid-base and electrolytes for sophomores, 1972-74

Specific administrative responsibilities

Assistant Director and Assistant Attending Pathologist, Division of Clinical Pathology, Upstate Medical Center, 1965-68
Hospital Emergency Room - Outpatient Committee, 1965-68
Hospital Safety Committee, 1965-68; Chairman, 1967-68
Syracuse Red Cross Blood Program Committee, 1966-68
Hospital Disaster Planning Committee, 1967-68
House Officers and Graduate Students Committee, 1967-68
Syracuse Red Cross Blood Recruitment Committee, 1967-68
Medical Occupations Advisory Committee, Manpower Development
Training Program, 1967-68; Chairman, Subcommittee on
Laboratory Training

Cancer Subcommittee, Onondaga County Medical Society, 1967-68
Blood Bank Inspector, Northeast District, American Association
of Blood Banks, 1967-68

General Director and Coordinator, American Association of Blood
Banks Pre-convention Workshop on Blood Component Therapy,
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Workshop Committee, American Association of Blood Banks, 1967-68
Member, Medical Technology Workshops Committee, American Society
of Clinical Pathologists, 1969-71

Member, Ad Hoc Review Committee for Advanced Traineeships, Division of Allied Health Manpower, NIH, 1969

Panel Member, Interagency Board of U. S. Civil Service Examiners for Washington, D.C., 1969-70

Military service

U. S. Public Health Service, 1968-70

Curriculum Vitae
Page 4
Other employment or activity

Other employment or activity

Reagents Section, Laboratory of Blood and Blood Products, Division of Biologics Standards, National Clinical Pathologist, Clinical Chemistry Service, Clinical Repartment, Clinical Center, NIH, January - June, 1969 Assistant Section Chief, Clinical Chemistry Service, Clinical Pathology Department, Clinical Center, NIH, July 1969 - June, 1970 Assistant Director (1970 - 71), Director (1971 - 74), Bio-Science Laboratories, Van Nuys, California in Professor and Director, Program in Pathology, University of Texas Medical School at Houston, Houston, Texas (1974 - Present)

Society Memberships:

D. Society Memberships: National Paragraphics

None

Research Activities:

Complete Bibliography Appended.

Major Areas of Research Interest

Clinical chemistry, especially correlation of clinical laboratory tests with diagnosis. The grant of a fixed and a land of

Research in Progress

Correlation of serum triidothyronine assays with thyroid status.

Research grants in Past Five Years

None

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BIBLIOGRAPHY

- 1. Cannon, D.C., and R. W. Wissler. Migration of Spleen Cells into the Blood Stream Following Antigen Stimulation of the Rat. Nature 207:654, 1965.
- 2. Ward, P.H., Cannon, D.C., and J.R. Lindsay. The Vestibular System in Multiple Sclerosis. A Clinical Histopathological Study.

 Laryngoscope 75:1031, 1965.
- 3. Cannon, D.C., and R.W. Wissler. Restoration of the Immune Response by Circulating Lymphocytes. Arch. Path. 83:188, 1967.
- 4. Cannon, D.C., and R.W. Wissler. Spleen Cell Migration in the Immune Response of the Rat. Arch. Path. 84:109, 1967.
- 5. Cannon, D.C., Immunoglobulin Analysis in Clinical Diagnosis.

 I. Quantitative Methods. Postgrad. Med. 46(2):55, 1969.
- 6. Cannon, D.C., Immunoglobulin Analysis in Clinical Diagnosis. II. Immunoelectrophoresis. Postgrad. Med. 46(3):55, 1969.
- 7. Cannon, D.C. Clinical Aspects of the Leukocyte Antibody Reaction.
 Postgrad. Med. 47:51, 1970.
- 8. Reed, A.H., Cannon, D.C., Winkelman, J.W., Bhasin, Y.P.,
 Henry, R.J., and V.J. Pileggi. Estimation of Normal Ranges from a
 Controlled Sample Survey. I. Sex and Age-Related Influence on the
 SMA-12/60 Screening Group of Tests. Clin. Chem. 18:57, 1972.
- 9. Reed, A.H., Cannon, D.C., Pileggi, V.J., and J.W. Winkelman. Use of confidence intervals to assess precision of normal range estimates. Clin. Biochem. 6:29, 1973.
- 10. Winkelman, J.W., Cannon, D.C., Pileggi, V.J., and A.H. Reed.

 Estimation of norms from a controlled sample survey. II.

 Influence of body habitus, oral contraceptives and other factors on values for the normal range derived from the SMA 12/60 screening group of tests. Clin. Chem. 19:488, 1973.
 - 11. Wybenga, D.R., Ibbott, F.A., and D.C. Cannon. Ionized calcium:
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